

initialseq.ST25.txt SEQUENCE LISTING

<110> Wang, Tongwen

<120> Composistions and Methods of modulating TGF-B Signaling

<130> 17633/1082

<140> 09927738

<141> 2001-08-10

<150> US 60/119786

<151> 1999-02-11

<150> PCT/US00/03561

<151> 2000-02-11

<160> 28

<170> PatentIn version 3.1

<210> 1

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<212> PRT

<213> Unknown

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<223> Isolated using yeast two hybrid system, Clone S1 + 27

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Ala Glu Lys Leu Ala Ala Ala Gln Gly Gln Ala Pro Leu Glu Pro Thr 35 40 45

Gln Asp Gly Ser Ala Ile Glu Thr Cys Pro Lys Gly Asp Glu Pro Arg 50 60

Gly Asp Glu Gln Gln Val Glu Ser Met Thr Pro Lys Pro Val Leu Gln 65 70 75

Glu Glu Asn Asn Gln Glu Ser Phe Ile Ala Phe Ala Arg Val Phe Ser 85 90 95

Gly Val Ala Arg Arg Gly Lys Lys Ile Phe Val Leu Gly Pro Lys Tyr 100 105 110

Ser Pro Leu Glu Phe Leu Arg Arg Val Pro Leu Cys Phe Ser Ala Pro 115 120 125

Pro Asp Gly Leu Pro Gln Val Pro His Met Ala Tyr Cys Ala Leu Glu 130 135 140

Asn Leu Tyr Leu Leu Met Gly Arg Glu Leu Glu Tyr Leu Glu Glu Val 145 150 150

Pro Pro Gly Asn Val Leu Gly Ile Gly Gly Leu Gln Asp Phe Val Leu 165 170 175

Page 3

Lys Ser Ala Thr Leu Cys Ser Leu Pro Ser Cys Pro Pro Phe Ile Pro 180 185 190

Leu Asn Phe Glu Ala Thr Pro Ile Val Arg Val Ala Val Glu Pro Lys 195 200 205

His Pro Ser Glu Met Pro Gln Leu Val Lys Gly Met Lys Leu Leu Asn 210 215 220

Gln Ala Asp Pro Cys Val Gln Ile Leu Ile Gln Glu Thr Gly Glu His 225 230 235 240

Val Leu Val Thr Ala Gly Glu Val His Leu Gln Arg Cys Leu Asp Asp 245 250 255

Leu Lys Glu Arg Phe Ala Lys Ile His Ile Ser Val Ser Glu Pro Ile 260 265 270

ile Pro Phe Arg Glu Thr Ile Thr Lys Pro Pro Lys Val Asp Met Val 275 280 285

Asn Glu Glu Ile Gly Lys Gln Gln Lys Val Ala Val Ile His Gln Met 290 295 300

Lys Glu Asp Gln Ser Lys Ile Pro Glu Gly Ile Gln Val Asp Ser Asp 305 310 315

Gly Leu Ile Thr Ile Thr Thr Pro Asn Lys Leu Ala Thr Leu Ser Val

Arg Ala Met Pro Leu Pro Glu Glu Val Thr Gln Ile Leu Glu Glu Asn 340 350

Ser Asp Leu Ile Arg Ser Met Glu Gln Leu Thr Ser Ser Leu Asn Glu 355 360 365

Gly Glu Asn Thr His Met Ile His Gln Lys Thr Gln Glu Lys Ile Trp 370 375 380

Glu Phe Lys Gly Lys Leu Glu Gln His Leu Thr Gly Arg Arg Trp Arg 385 390 395

Asn Ile Val Asp Gln Ile Trp Ser Phe Gly Pro Arg Lys Cys Gly Pro 415

Asn Ile Leu Val Asn Lys Ser Glu Asp Phe Gln Asn Ser Val Trp Thr 420 425 430 Page 4

Gly Pro Ala Asp Lys Ala Ser Lys Glu Ala Ser Arg Tyr Arg Asp Leu 435 440 445

Gly Asn Ser Ile Val Ser Gly Phe Gln Leu Ala Thr Leu Ser Gly Pro 450 455 460

Met Cys Glu Glu Pro Leu Met Gly Val Cys Phe Val Leu Glu Lys Trp 465 470 475 480

Asp Leu Ser Lys Phe Glu Glu Gln Gly Ala Ser Asp Leu Ala Lys Glu 485 490 495

Asp Arg Arg Lys Met Lys Pro Val Leu Val Glu Met Lys Thr Lys Ser 500 505

Tyr Lys Met Ala Ala Leu Arg Pro Leu Arg Arg Gly His His Arg Lys 515 520 525

Glu Asn Leu His Ser Leu Thr Ala Met Asp Leu Ser Gln Asp Ser 530 535 540

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Ser Gly Gly Ser Pro Ser Pro Pro Thr Ser Glu Pro Ala Arg Ser Gly 50 60

His Arg Gly Asn Arg Ala Arg Gly Val Ser Arg Ser Pro Pro Lys Lys Page 5 70

Lys Asn Lys Ala Ser Gly Arg Arg Ser Lys Ser Pro Arg Ser Lys Arg 85 90 95 Asn Arg Ser Pro His His Ser Thr Val Lys Val Lys Gln Glu Arg Glu 100 105 110 Asp His Pro Arg Arg Gly Arg Glu Asp Arg Gln His Arg Glu Pro Ser 115 120 125 Glu Gln Glu His Arg Arg Ala Arg Asn Ser Asp Arg Asp Arg His Arg 130 135 140 Gly His Ser His Gln Arg Arg Thr Ser Asn Glu Arg Pro Gly Ser Gly 145 150 150 160 Gln Gly Gln Gly Arg Asp Arg Asp Thr Gln Asn Leu Gln Ala Gln Glu 165 170 175 Glu Glu Arg Glu Phe Tyr Asn Ala Arg Arg Arg Glu His Arg Gln Arg 180 185 190 Asn Asp Val Gly Gly Gly Ser Glu Ser Gln Glu Leu Val Pro Arg 195 200 205 Pro Gly Gly Asn Asn Lys Glu Lys Glu Val Pro Ala Lys Glu Lys Pro 210 215 220 Ser Phe Glu Leu Ser Gly Ala Leu Leu Glu Asp Thr Asn Thr Phe Arg 225 230 235 240 Gly Val Val Ile Lys Tyr Ser Glu Pro Pro Glu Ala Arg Ile Pro Lys 245 250 255 Lys Arg Trp Arg Leu Tyr Pro Phe Lys Asn Asp Glu Val Leu Pro Val 260 265 270 Met Tyr Ile His Arg Gln Ser Ala Tyr Leu Leu Gly Arg His Arg Arg 275 280 285 Ile Ala Asp Ile Pro Ile Asp His Pro Ser Cys Ser Lys Gln His Ala 290 295 300 Val Phe Gln Tyr Arg Leu Val Glu Tyr Thr Arg Ala Asp Gly Thr Val 305 310 315

Page 6

initialseq.ST25.txt Gly Arg Arg Val Lys Pro Tyr Ile Ile Asp Leu Gly Ser Gly Asn Gly 325 330 335 Thr Phe Leu Asn Asn Lys Arg Ile Glu Pro Gln Arg Tyr Tyr Glu Leu 340 345 350 Lys Glu Lys Asp Val Leu Lys Phe Gly Phe Ser Ser Arg Glu Tyr Val 355 360 365 Leu Leu His Glu Ser Ser Asp Thr Ser Glu Ile Asp Arg Lys Asp Asp 370 380 Glu Asp Glu Glu Glu Glu Glu Val Ser Asp Ser 385 390 395 <210> <211> <212> PRT <213> Unknown <220> <223> Protein Sequence of NIPP-1 domain homologous to SNIP 1 <400> Tyr Leu Phe Gly Arg Asn Pro Asp Leu Cys Asp Phe Thr Ile Asp His
1 10 15 Gln Ser Cys Ser Arg Val His Ala Ala Leu Val Tyr His Lys His Leu 20 25 30 Lys Arg Val Phe Leu Ile Asp Leu Asn Ser Thr His Gly Thr Phe Leu 35 40 45 Gly His Ile Arg Leu Glu Pro His Lys Pro Gln Gln Ile Pro Ile Asp 50 55 60 Ser Thr Val Ser Phe Gly Ala Ser Thr Arg Ala Tyr Thr Leu Arg Glu 65 70 75 80 Lys Pro

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Ala Gln Glu Glu Arg Glu Phe Tyr Asn Ala Arg Arg Glu His
35 40 45

Arg Gln Arg Asn Asp Val Gly Gly Gly Gly Ser Glu Ser Gln Glu Leu 50 55 60

Val Pro Arg Pro Gly Gly Asn Asn Lys Glu Lys Glu Val Pro Ala Lys 65 70 75 80

Glu Lys Pro Ser Phe Glu Leu Ser Gly Ala Leu Leu Glu Asp Thr Asn 85 90 95

Thr Phe Arg Gly Val Val Ile Lys Tyr Ser Glu Pro Pro Glu Ala Arg 100 105 110

Ile Pro Lys Lys Arg Trp Arg Leu Tyr Pro Phe Lys Asn Asp Glu Val 115 120 125

Leu Pro Val Met Tyr Ile His Arg Gln Ser Ala Tyr Leu Leu Gly Arg 130 135 140

His Arg Arg Ile Ala Asp Ile Pro Ile Asp His Pro Ser Cys Ser Lys 145 150 160

Gln His Ala Val Phe Gln Tyr Arg Leu Val Glu Tyr Thr Arg Ala Asp 165 170 175

Gly Thr Val Gly Arg Arg Val Lys Pro Tyr Ile Ile Asp Leu Gly Ser 180 185 190

Gly Asn Gly Thr Phe Leu Asn Asn Lys Arg Ile Glu Pro Gln Arg Tyr 195 200 205 Page 8

Tyr Glu Leu Lys Glu Lys Asp Val Leu Lys Phe Gly Phe Ser Ser Arg 210 215 220

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Tyr Pro Phe Lys Gly Glu Glu Ser Leu Gln Val Leu Tyr Ile His Arg 35 40 45

Gln Ser Ala Tyr Leu Ile Gly Arg Asp His Lys Ile Ala Asp Ile Pro 50 60

Val Asp His Pro Ser Cys Ser Lys Gln His Ala Val Leu Gln Phe Arg 65 70 75 80

Ser Met Pro Phe Thr Arg Asp Asp Gly Thr Lys Ala Arg Arg Ile Met 85 90 95

Pro Tyr Ile Ile Asp Leu Gly Ser Gly Asn Gly Thr Phe Leu Asn Glu 100 105 110

Lys Lys Ile Glu Pro Gln Arg Tyr Ile Glu Leu Gln Glu Lys Asp Met 115 120 125

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Gln Ser Asp Ala Ala Leu Gln Val Asp Ile Ser Asp Ala Leu Ser Glu 35 40 45

Arg Asp Lys Val Lys Phe Thr Val His Thr Lys Ser Ser Leu Pro Asn 50 55 60

Phe Lys Gln Asn Glu Phe Ser Val Val Arg Gln His Glu Glu Phe Ile 65 70 75 80

Trp Leu His Asp Ser Phe Val Glu Asn Glu Asp Tyr Ala Gly Tyr Ile 85 90 95

Ile Pro Pro Ala Pro Pro Arg Pro Asp Phe Asp Ala Ser Arg Glu Lys
100 105 110

Thr Lys Met Lys Gln Glu Leu Glu Ala Glu Tyr Leu Ala Ile Phe Lys 130 135 140 initialseq.ST25.txt Lys Thr Val Ala Met His Glu Val Phe Leu Cys Arg Val Ala Ala His 145 150 155 160 Pro Ile Leu Arg Arg Asp Leu Asn Phe His Val Phe Leu Glu Tyr Asn 165 170 175 Gln Asp Leu Ser Val Arg Gly Lys Lys Lys Lys Lys Asn Ser Arg Ser 180 185 190 Phe Gly Leu Leu Arg Gln 195 <210> 8 <211> 414 **PRT** <212> <213> Unknown <220> Clone S1+12 -2 protein Sequence <223> <400> His Ala Ser Gly Leu Gly Ala Ala Met Met Glu Gly Leu Asp Asp Gly $10 \ 15$ Pro Asp Phe Leu Ser Glu Glu Asp Arg Gly Leu Lys Ala Ile Asn Val 20 25 30 Asp Leu Gln Ser Asp Ala Ala Leu Gln Val Asp Ile Ser Asp Ala Leu 35 40 45 Ser Glu Arg Asp Lys Val Lys Phe Thr Val His Thr Lys Ser Ser Leu 50 60 Pro Asn Phe Lys Gln Asn Glu Phe Ser Val Val Arg Gln His Glu Glu 65 70 75 Phe Ile Trp Leu His Asp Ser Phe Val Glu Asn Glu Asp Tyr Ala Gly 85 90 95 Tyr Ile Ile Pro Pro Ala Pro Pro Arg Pro Asp Phe Asp Ala Ser Arg 100 105 110

Page 11

Glu Lys Leu Gln Lys Leu Gly Glu Gly Glu Gly Ser Met Thr Lys Glu 115 120 125

Glu Phe Thr Lys Met Lys Gln Glu Leu Glu Ala Glu Tyr Leu Ala Ile Phe Lys Lys Thr Val Ala Met His Glu Val Phe Leu Cys Arg Val Ala 145 150 155 160 Ala His Pro Ile Leu Arg Arg Asp Leu Asn Phe His Val Phe Leu Glu 165 170 Tyr Asn Gln Asp Leu Ser Val Arg Gly Lys Asn Lys Lys Glu Lys Leu 180 185 190 Glu Asp Phe Phe Lys Asn Met Val Lys Ser Ala Asp Gly Val Ile Val 195 200 205 Ser Gly Val Lys Asp Val Asp Asp Phe Phe Glu His Glu Arg Thr Phe 210 220 Leu Leu Glu Tyr His Asn Arg Val Lys Asp Ala Ser Ala Lys Ser Asp 225 230 235 240 Arg Met Thr Arg Ser His Lys Ser Ala Ala Asp Asp Tyr Asn Arg Ile 245 250 255 Gly Ser Ser Leu Tyr Ala Leu Gly Thr Gln Asp Ser Thr Asp Ile Cys 260 265 270 Lys Phe Phe Leu Lys Val Ser Glu Leu Phe Asp Lys Thr Arg Lys Ile 275 280 285 Glu Ala Arg Val Ser Ala Asp Glu Asp Leu Lys Leu Ser Asp Leu Leu 290 295 300 Lys Tyr Tyr Leu Arg Glu Ser Gln Ala Ala Lys Asp Leu Leu Tyr Arg 305 310 315 Arg Ser Arg Ser Leu Val Asp Tyr Glu Asn Ala Asn Lys Ala Leu Asp 325 330 335 Lys Ala Arg Ala Lys Asn Lys Asp Val Leu Gln Ala Glu Thr Ser Gln 340 345 350Gln Leu Cys Cys Gln Lys Phe Glu Lys Ile Ser Glu Ser Ala Lys Gln 355 360 365 Glu Leu Ile Asp Phe Lys Thr Arg Arg Val Ala Ala Phe Arg Lys Asn 370 380 Page 12

Leu Val Glu Leu Ala Glu Leu Glu Leu Lys His Ala Lys Gly Asn Leu 385 390 395 400

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Glu Val Phe Leu Cys Arg Val Ala Ala His Pro Ile Leu Arg Arg Asp 35 40 45

Leu Asn Phe His Val Phe Leu Glu Tyr Asn Gln Asp Leu Ser Val Arg 50 55 60

Gly Lys Asn Lys Lys Glu Lys Leu Glu Asp Phe Phe Lys Asn Met Val 65 70 80

Lys Ser Ala Asp Gly Val Ile Val Ser Gly Val Lys Asp Val Asp Asp 85 90 95

Phe Phe Glu His Glu Arg Thr Phe Leu Leu Glu Tyr His Asn Arg Val 100 105 110

Lys Asp Ala Ser Ala Lys Ser Asp Arg Met Thr Arg Ser His Lys Ser 115 120 125

Ala Ala Asp Asp Tyr Asn Arg Ile Gly Ser Ser Leu Tyr Ala Leu Gly 130 135 140

Thr Gln Asp Ser Thr Asp Ile Cys Lys Phe Phe Leu Lys Val Ser Glu Page 13

160

Leu Phe Asp Lys Thr Arg Lys Ile Glu Ala Arg Val Ser Ala Asp Glu 165 170 175

Asp Leu Lys Eeu Ser Asp Leu Leu Lys Tyr Tyr Leu Arg Glu Ser Gln 180 185 190

Ala Ala Lys Asp Leu Leu Tyr Arg Arg Ser Arg Ser Leu Val Asp Tyr 195 200 205

Glu Asn Ala Asn Lys Ala Leu Asp Lys Ala Arg Ala Lys Asn Lys Asp 210 220

Val Leu Gln Ala Glu Thr Ser Gln Gln Leu Cys Cys Gln Lys Phe Glu 225 230 235 240

Lys Ile Ser Glu Ser Ala Lys Gln Glu Leu Ile Asp Phe Lys Thr Arg 245 250 255

Arg Val Ala Ala Phe Arg Lys Asn Leu Val Glu Leu Ala Glu Leu Glu 260 265 270

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<213> Unknown

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ggacttcaga	gagaatgtat	tttccagttt	actggcaatt	ttgcgtcatt	ttgttgggac	240
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Leu	Lys	Cys 35	Cys	val	Cys	Lys	Lys 40	Asn	Gly	Ala	Ser	11e 45	Gly	Cys	val
Αla	Pro 50	Arg	Cys	Lys	Arg	ser 55	Tyr	His	Phe	Pro	Cys 60	Gly	Leu	Gln	Arg
Glu 65	Cys	Ile	Phe	Gln	Phe 70	Thr	Glу	Asn	Phe	а1а 75	Ser	Phe	Cys	Trp	Asp 80
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11e		Asr	ı Asn	Ser	Asp 150	Ile	Phe	Gln	Lys	Glu 155	Met	Leu	Arg	Met	Gly 160
Ιle	e His	: Ile	e Pro	Glu 165	Lys	Asp	Ala	Ser	Trp 170	Glu	Leu	Glu	Glu	Asn 175	Ala
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Ar	g Cys	5 Lys 19	s Glu 5	ı Gly	/ Arg	ı Asp	туг 200	Asn	ıAla	Pro	Asp	Ser 205	Lys	тгр	Glu
Ιl	e Ly:	s Ar	g Cy:	s Glr	ı Cys	Cys 215	Gly	' Ser	· Ser	·Gly	7 Thr 220	His	. Lei	ıAla	Cys
Se 22		r Le	u Ar	g Se	r Trp 230	o Glu	ı Glr	n Asr	ı Trp	o Glu 235	ı Cys	Lei	ı Glu	ı Cys	240
G٦	y Il	e Il	е ту	r Ası 24	n Sei 5	r Gly	y Glu	ı Phe	e Glr 250	n Thi	ala	Ly:	s Ly:	s His 255	val

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Ser Ser Pro Lys Leu Pro Arg Gln Ser Pro Gly Ser Gln Ser Lys Asp 275 280 285

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Thr Gly Ile Gly His Gln Thr Ile Pro Val Ser Leu Pro Ala Ala Gly 50 60

Met Gly His Gln Ala Arg Gly Met Ser Leu Gln Ser Asn Tyr Leu Gly Page 18 Leu Ala Ala Ala Pro Ala Ile Met Ser Tyr Ala Glu Cys Ser Val Pro 85 90 95

Ile Gly Val Thr Ala Pro Ser Leu Gln Pro Val Gln Ala Arg Gly Ala 100 105 110

Val Pro Thr Ala Thr Ile Ile Glu Pro Pro Pro Pro Pro Pro Pro 115 120 125

Pro Pro Pro Pro Pro Ala Pro Lys Met Pro Pro Glu Lys Thr 130 140

Lys Lys Gly Arg Lys Asp Lys Ala Lys Lys Ser Lys Thr Lys Met Pro 145 155 160

Ser Leu Val Lys Lys Trp Gln Ser Ile Gln Arg Glu Leu Asp Glu Glu 165 170 175

Asp Asn Ser Ser Ser Glu Glu Asp Arg Glu Ser Thr Ala Gln Lys 180 185 190

Arg Ile Glu Glu Trp Lys Gln Gln Gln Leu Val Ser Gly Met Ala Glu 195 200 205

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Page 19

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Pro Gly Phe Gly Val Asp Gln Leu Arg Asp Asp Asn Leu Glu Thr Tyr 85 90 95

Trp Gln Ser Asp Gly Ser Gln Pro His Leu Val Asn Ile Gln Phe Arg

Arg Lys Thr Thr Val Lys Thr Leu Cys Ile Tyr Ala Asp Tyr Lys Ser 115 120 125

Asp Glu Ser Tyr Thr Pro Ser Lys Ile Ser Val Arg Val Gly Asn Asn 130 135 140

Phe His Asn Leu Gln Glu Ile Arg Gln Leu Glu Leu Val Glu Pro Ser 145 150 160

Gly Trp Ile His Val Pro Leu Thr Asp Asn His Lys Lys Pro Thr Arg 165 170 175

Thr Phe Met Ile Gln Ile Ala Val Leu Ala Asn His Gln Asn Gly Arg 180 185 190

Asp Thr His Met Arg Gln Ile Lys Ile Tyr Thr Pro Val Glu Glu Ser 195 200 205

Ser Ile Gly Lys Phe Pro Arg Cys Thr Thr Ile Asp Phe Met Met Tyr 210 220

Arg Ser Ile Arg Leu Asn Glu Thr Lys Ile Ile Lys Arg Ile Phe Val 225 230 235

Leu Ile Leu Tyr Leu Asn Asn Ile Ser Cys Thr Phe Ile Glu Gln Gly 245 250 255

Ile Arg Tyr Ile Phe Cys Ile Cys Leu Lys Ile Phe Tyr Cys Asn Phe 260 265 270

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Page 21

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Gly Leu Gln Ala Gly Leu Pro Gly Phe Tyr Asp Pro Cys Val Gly Glu 65 70 75 80	
Glu Lys Asn Leu Lys Val Leu Tyr Gln Phe Arg Gly Val Leu His Gln 85 90 95	
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